

```
g nodes:

1 2 3 4 5 6 7 8 9
in bonds:

1-11 1-12 3-13 4-14 9-10 12-16 13-15
g bonds:

1-2 1-5 2-3 3-4 4-5 4-6 5-9 6-7 7-8 8-9
ct/norm bonds:

1-2 1-5 1-12 2-3 3-4 3-13 4-5 4-6 4-14 5-9 6-7 7-8 8-9 9-10
ct bonds:

1-11 12-16 13-15
```

in nodes : ' 10 11 12 13 14 15 16

ch level : 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS Welcome to STN International! Enter x:x

LOGINID:ssspta1623kxg PASSWORD: TERMINAL (ENTER 1, 2, 3, OR ?):2 Welcome to STN International Web Page URLs for STN Seminar Schedule - N. America NEWS "Ask CAS" for self-help around the clock NEWS 2 NEWS SEP 09 CA/CAplus records now contain indexing from 1907 to the present NEWS AUG 05 New pricing for EUROPATFULL and PCTFULL effective August 1, 2003 NEWS 5 AUG 13 Field Availability (/FA) field enhanced in BEILSTEIN NEWS 6 AUG 18 Data available for download as a PDF in RDISCLOSURE NEWS 7 AUG 18 Simultaneous left and right truncation added to PASCAL NEWS 8 AUG 18 FROSTI and KOSMET enhanced with Simultaneous Left and Righ Truncation NEWS 9 AUG 18 Simultaneous left and right truncation added to ANABSTR NEWS 10 SEP 22 DIPPR file reloaded DEC 08 NEWS 11 INPADOC: Legal Status data reloaded SEP 29 NEWS 12 DISSABS now available on STN OCT 10 NEWS 13 PCTFULL: Two new display fields added OCT 21 BIOSIS file reloaded and enhanced NEWS 14 BIOSIS file segment of TOXCENTER reloaded and enhanced NEWS 15 OCT 28 NEWS 16 NOV 24 MSDS-CCOHS file reloaded DEC 08 NEWS 17 CABA reloaded with left truncation NEWS 18 DEC 08 IMS file names changed NEWS 19 DEC 09 Experimental property data collected by CAS now available in REGISTRY NEWS 20 DEC 09 STN Entry Date available for display in REGISTRY and CA/CAplus NEWS 21 DEC 17 DGENE: Two new display fields added NEWS 22 DEC 18 BIOTECHNO no longer updated NEWS 23 DEC 19 CROPU no longer updated; subscriber discount no longer available DEC 22 Additional INPI reactions and pre-1907 documents added to CAS NEWS 24 databases NEWS 25 DEC 22 IFIPAT/IFIUDB/IFICDB reloaded with new data and search fields NEWS EXPRESS NOVEMBER 14 CURRENT WINDOWS VERSION IS V6.01c, CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP), AND CURRENT DISCOVER FILE IS DATED 23 SEPTEMBER 2003 NEWS HOURS STN Operating Hours Plus Help Desk Availability NEWS INTER General Internet Information NEWS LOGIN Welcome Banner and News Items Direct Dial and Telecommunication Network Access to STN NEWS PHONE

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CAS World Wide Web Site (general information)

* * * * * * * * * * * * * * STN Columbus * * * * * * * * * * * * *

FILE 'HOME' ENTERED AT 10:53:49 ON 22 DEC 2003

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 10:53:59 ON 22 DEC 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

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STRUCTURE FILE UPDATES: 21 DEC 2003 HIGHEST RN 629597-20-2 DICTIONARY FILE UPDATES: 21 DEC 2003 HIGHEST RN 629597-20-2

TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2003

Please note that search-term pricing does apply when conducting ${\tt SmartSELECT}$ searches.

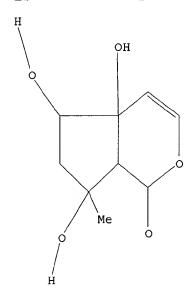
Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=> Uploading 09995691-3.str

L1 STRUCTURE UPLOADED

=> d l1 L1 HAS NO ANSWERS L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 11 sss sam SAMPLE SEARCH INITIATED 10:54:37 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 853 TO ITERATE

100.0% PROCESSED 853 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS:

15308 TO 18812

PROJECTED ANSWERS:

2 TO 124

1110020122 12111212

2 10

L2

2 SEA SSS SAM L1

=> d scan

L2 2 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

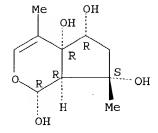
IN Cyclopenta[c]pyran-4-carboxylic acid, 1-(.beta.-D-glucopyranosyloxy)1,4a,5,6,7,7a-hexahydro-4a,5,6,7-tetrahydroxy-7-methyl-, methyl ester,
(1S,4aR,5R,6S,7R,7aR)- (9CI)

MF C17 H26 O13

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):1



ALL ANSWERS HAVE BEEN SCANNED

=> s l1 sss full FULL SEARCH INITIATED 10:55:07 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 17637 TO ITERATE

100.0% PROCESSED 17637 ITERATIONS

33 ANSWERS

SEARCH TIME: 00.00.02

L3 33 SEA SSS FUL L1

=> file caplus
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST

148.55 148.76

FILE 'CAPLUS' ENTERED AT 10:55:20 ON 22 DEC 2003
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FILE COVERS 1907 - 22 Dec 2003 VOL 139 ISS 26 FILE LAST UPDATED: 21 Dec 2003 (20031221/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 13 and (arthritis or osteoporosis or disc)

165 L3

31062 ARTHRITIS

2 ARTHRITISES

31062 ARTHRITIS

(ARTHRITIS OR ARTHRITISES)

12642 OSTEOPOROSIS

12163 DISC

2833 DISCS

14639 DISC

(DISC OR DISCS)

4 L3 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)

=> dis 14 1-4 bib abs hitstr

L4 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN

AN 2002:799804 CAPLUS

DN 138:103853

TI In vitro propagation and iridoid analysis of the medicinal species Harpagophytum procumbens and H. zeyheri

AU Levieille, G.; Wilson, G.

CS Department of Botany, University College Dublin, Dublin, Ire.

Plant Cell Reports (2002), 21(3), 220-225

CODEN: PCRPD8; ISSN: 0721-7714

PB Springer-Verlag

DT Journal

SO

AB

TΤ

CN

LA English

Exts. of the tubers of Harpagophytum procumbens DC (Devil's Claw) are used widely for the relief of arthritis, lumbago and muscular pain. The anti-inflammatory activity has been attributed to their iridoid components. A two-step protocol was established for the in vitro propagation of plants of Harpagophytum sp. by the regeneration of new plantlets from nodal cuttings and their acclimatization to ex vitro conditions. Single node cuttings were submitted to a root induction treatment with .beta.-indoleacetic acid (5 days at 2 mg l-1) followed by a transfer to a phytohormone-free medium to promote root elongation and support plantlet development. The new plantlets were weaned under autotrophic conditions and subsequently acclimatized in a glasshouse where they grew into fertile flowering plants that produced the characteristic Devil's Claw fruits as well as tuberised roots. Anal. of the tuber tissue of the micropropagated plants showed the presence of the iridoids harpagoside and harpagide at concns. comparable with those found in the wild plant material (1% dry wt.). The leaves were also found to contain these iridoids, and therefore could potentially provide an alternative and more sustainable source of therapeutically active compds. The application of in vitro methods for the propagation of Devil's Claw would contribute to the conservation of this species.

6926-08-5, Harpagide

RL: NPO (Natural product occurrence); BIOL (Biological study); OCCU (Occurrence)

(in vitro propagation and iridoid anal. of medicinal species Harpagophytum procumbens and H. zeyheri)

RN 6926-08-5 CAPLUS

.beta.-D-Glucopyranoside, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI) (CA INDEX NAME)

RE.CNT 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L4 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN
- AN 2002:591953 CAPLUS
- DN 137:159305
- TI 2-0-(9Z,12Z-octadecadienoyl)-3-0-[.alpha.-D-galactopyranosyl-(1''-6')-0-.beta.-D-galactopyranosyl]glycerol and pharmaceutical formulations containing it
- IN Shin, Jun Shik; Kim, Sang Tae; Hahn, Yong Nam
- PA Japan
- SO Jpn. Kokai Tokkyo Koho, 30 pp.
 - CODEN: JKXXAF
- DT Patent
- LA Japanese
- FAN.CNT 1

| T NO. KIND DATE | | APPLICATION NO. | DATE | | |
|-----------------|----------|----------------------------|--|--|--|
| | | | | | |
| A2 | 20020809 | JP 2001-365399 | 20011129 | | |
| B1 | 20030311 | US 2001-995617 | 20011129 | | |
| Α | 20001129 | | | | |
| | A2
B1 | A2 20020809
B1 20030311 | A2 20020809 JP 2001-365399
B1 20030311 US 2001-995617 | | |

Pharmaceutical formulations for treatment of osteoporosis,
arthritis, or intervertebral disk hernia, contain
2-0-(9Z,12Z-octadecadienoyl)-3-0-[.alpha.-D-galactopyranosyl-(1''-6')-0.beta.-D-galactopyranosyl]glycerol (I) or its esters as active
ingredients. I (420 mg) was purified from an EtOH ext. of 1848 g Cibotium
barometz root powder. Administration of I at 75 .mu.g/mL p.o. for 2 wk

prevented mouse paw edema induced by Zymosan A and Freund's adjuvant. Formulation examples of injections, tablets, capsules, and liqs. contg. I or I acetate are given.

IT 6926-08-5, Harpagide

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceuticals contg. octadecadienoyl(galactopyranosylgalactopyranosy

- 1) glycerol for treatment of osteoporosis, arthritis
- , and intervertebral disk hernia)
- RN 6926-08-5 CAPLUS
- CN .beta.-D-Glucopyranoside, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI) (CA INDEX NAME)

- L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN
- AN 2002:533182 CAPLUS
- DN 137:88448
- TI Use of harpaqide-related compounds for prevention and treatment of

osteoporosis, arthritis, and intervertebral disk hernia, pharmaceutical compositions, and preparation of the compounds Shin, Jun Sik; Kim, Sang-Tae; Hahn, Yong-Nam

IN PΑ S. Korea

Jpn. Kokai Tokkyo Koho, 29 pp. SO

CODEN: JKXXAF

DTPatent

Japanese LA

| FAN.CNT 1 | | | | |
|---------------------|------|----------|-----------------|----------|
| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
| | | | | |
| PI JP 2002201136 | A2 | 20020716 | JP 2001-365400 | 20011129 |
| US 2002183264 | A1 | 20021205 | US 2001-995691 | 20011129 |
| PRAI KR 2000-71497 | A | 20001129 | | |
| OS MARPAT 137:88448 | } | | | |
| GT | | | | |

Harpaqide-related compds. I (R1 = H, lower alkyl; R2 = H, cinnamoyl) are AΒ used for treatment or prevention of osteoporosis, arthritis, and/or intervertebral disk diseases. I (R1 = H, lower alkyl; R2 = cinnamoyl) are hydrolyzed to give I (R1 = H, lower alkyl; R2 = H). Harpagide (purified from Harpagophytum procumbens root) (at 75 .mu.g/kg/day p.o.) inhibited zymosan A- and Freund's adjuvant-induced rat paw edema. Formulation examples of injections, tablets, capsules, and ligs. contg. harpagide or harpagoside are given.

6926-08-5P, Harpagide IT

RL: PAC (Pharmacological activity); PUR (Purification or recovery); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(harpagide-related compds. for prevention and treatment of osteoporosis, arthritis, and intervertebral disk hernia)

6926-08-5 CAPLUS RN

.beta.-D-Glucopyranoside, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI) (CA INDEX NAME)

```
ANSWER 4 OF 4 CAPLUS COPYRIGHT 2003 ACS on STN
L4
      2001:31340 CAPLUS
AN
DN
      134:95502
      Compositions and methods for treating or preventing osteoporosis
TΙ
      Prince, Richard Lewis; Min, Xu
IN
      University of Western Australia, Australia; Guangzhou University of
PΑ
      Traditional Chinese Medicine
      PCT Int. Appl., 93 pp.
SO
      CODEN: PIXXD2
DT
      Patent
      English
LA
FAN.CNT 1
      PATENT NO.
                         KIND DATE
                                                 APPLICATION NO.
                                                                     DATE
                                                  ____
                                _____
                         A1
                                                 WO 2000-AU737
                                                                     20000629
      WO 2001001996
                                20010111
PΙ
      WO 2001001996
                         C2
                                20020912
              AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
               CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
               HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
               SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                19990629
PRAI AU 1999-1273
                         Α
     The invention relates to a therapeutic compn. and method for treating
      osteoporosis and other calcium, and/or estrogen related disorders.
      Examples are given for treating osteoporosis with exts. of
      plants such as Epimedium koreanum, Slavia miltiorrhiza, Asragalus
     membranaceus, Pueraria thomsonii, and Psoralea coryliofolia.
     6926-08-5, Harpagide
ΙT
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); THU
      (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)
         (herb medicine exts. for treating or preventing osteoporosis)
RN
      6926-08-5 CAPLUS
      .beta.-D-Glucopyranoside, (1S, 4aS, 5R, 7S, 7aR) -1, 4a, 5, 6, 7, 7a-hexahydro-
CN
      4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI) (CA INDEX NAME)
```

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

TOTAL COST IN U.S. DOLLARS SINCE FILE SESSION ENTRY 173.32 FULL ESTIMATED COST 24.56 TOTAL SINCE FILE DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SESSION ENTRY -2.60 CA SUBSCRIBER PRICE -2.60

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STRUCTURE FILE UPDATES: 21 DEC 2003 HIGHEST RN 629597-20-2 DICTIONARY FILE UPDATES: 21 DEC 2003 HIGHEST RN 629597-20-2

TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=> s 6926-08-5 L5 1 6926-08-5 (6926-08-5/RN)

=> s 15 and (arthritis or osteoporosis or disc)

273 ARTHRITIS 0 OSTEOPOROSIS

222 DISC

18 DISCS

239 DISC

(DISC OR DISCS)

L6 0 L5 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)

=> d scan 13

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-4-carboxylic acid, 6-chloro-1-(.beta.-D-glucopyranosyloxy)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methyl-, methyl ester, (1S,4aR,5S,6R,7R,7aS)- (9CI)

MF C17 H25 Cl O12

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):2-33 $^{\prime}$ 2-33 $^{\prime}$ IS NOT VALID HERE

To display more answers, enter the number of answers you would like to see. To end the display, enter "NONE", "N", "0", or "END". HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):5

- L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN
- IN .beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7methylcyclopenta[c]pyran-1-yl, 2,3,4,6,?,?-hexaacetate,
 [1S-(1.alpha.,4a.alpha.,5.alpha.,7.alpha.,7a.alpha.)]- (9CI)
- MF C27 H36 O16
- CI IDS

CM 1

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-4-carboxylic acid, 6-chloro-1-(.beta.-Dglucopyranosyloxy)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methyl-,
methyl ester, (1S,4aR,5S,6S,7R,7aS)- (9CI)
MF C17 H25 Cl O12

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-4-carboxylic acid, 1-(.beta.-D-glucopyranosyloxy)1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methyl-, methyl ester,
(1S,4aR,5R,7S,7aR)- (9CI)

MF C17 H26 O12

Absolute stereochemistry. Rotation (-).

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, (1S, 4aR, 5R, 7S, 7aR) -1, 4a, 5, 6, 7, 7a-hexahydro-

4a,5,7-trihydroxy-4,7-dimethylcyclopenta[c]pyran-1-yl (9CI)

MF C16 H26 O10

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, (1S, 4aS, 5R, 6S, 7S, 7aR) -1, 4a, 5, 6, 7, 7a-hexahydro-

4a,5,6,7-tetrahydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

MF C15 H24 O11

Absolute stereochemistry. Rotation (-).

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):5

REGISTRY COPYRIGHT 2003 ACS on STN L3 33 ANSWERS

Cyclopenta[c]pyran-1,4a,5,7(1H)-tetrol, 5,6,7,7a-tetrahydro-4,7-dimethyl-,
[1S-(1.alpha.,4a.beta.,5.beta.,7.beta.,7a.beta.)]- (9CI) IN

MF C10 H16 05

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

REGISTRY COPYRIGHT 2003 ACS on STN L3 33 ANSWERS

IN Propanoic acid, 3-[[(1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7trihydroxy-7-methylcyclopenta[c]pyran-1-yl]oxy]-2-hydroxy-3-[1-

(hydroxymethyl) -2-methoxy-2-oxoethoxy] -, methyl ester (9CI)

MF C17 H26 O12

Absolute stereochemistry. Currently available stereo shown.

REGISTRY COPYRIGHT 2003 ACS on STN L333 ANSWERS

.beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-IN methylcyclopenta[c]pyran-1-yl 6-0-.alpha.-D-glucopyranosyl-, [1S-(1.alpha.,4a.alpha.,5.alpha.,7.alpha.,7a.alpha.)]- (9CI) C21 H34 O15 MF

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, (1S,4aS,5S,6R,7R,7aS)-6-chloro-1,4a,5,6,7,7ahexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

MF C15 H23 Cl O10

Absolute stereochemistry. Rotation (-).

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, (1S, 4aS, 5R, 6S, 7R, 7aR) -1, 4a, 5, 6, 7, 7a-hexahydro-

4a,5,6,7-tetrahydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

MF C15 H24 O11

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):10

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-1,4a,5,7(1H)-tetrol, 5,6,7,7a-tetrahydro-4,7-dimethyl-,

[1R-(1.alpha.,4a.alpha.,5.alpha.,7.alpha.,7a.alpha.)]- (9CI)

MF C10 H16 O5

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

.beta.-D-Glucopyranoside, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl, 6-[(2E)-3-(4methoxyphenyl)-2-propenoate] (9CI)

MF C25 H32 O12

Absolute stereochemistry. Rotation (-). Double bond geometry as shown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7methylcyclopenta[c]pyran-1-yl, 6-[3-(4-hydroxyphenyl)-2-propenoate],
 [1S-[1.alpha.(E),4a.alpha.,5.alpha.,7a.alpha.]]- (9CI)

MF C24 H30 O12

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,6,7-tetrahydroxy-7methylcyclopenta[c]pyran-1-yl, [1S-(1.alpha.,4a.alpha.,5.beta.,6.beta.,7.a
lpha.,7a.alpha.)]- (9CI)

MF C15 H24 O11

CI COM

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-4-carboxylic acid, 1-(.beta.-D-glucopyranosyloxy)1,4a,5,6,7,7a-hexahydro-4a,5,6,7-tetrahydroxy-7-methyl-, methyl ester,
(1R,4aR,5R,6S,7R,7aR)- (9CI)

MF C17 H26 O13

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

.beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl, monoacetate, [1S-

(1.alpha., 4a.alpha., 5.alpha., 7.alpha., 7a.alpha.)] - (9CI)

MF C17 H26 O11

CI IDS

CM 1

Absolute stereochemistry.

CM 2

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl, 6-[(2Z)-3-(4-

methoxyphenyl)-2-propenoate] (9CI)
MF C25 H32 O12

Absolute stereochemistry. Rotation (-). Double bond geometry as shown.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

.beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7methylcyclopenta[c]pyran-1-yl, 6-[3-(4-hydroxyphenyl)-2-propenoate],
 [1s-[1.alpha.(Z),4a.alpha.,5.alpha.,7a.alpha.]]- (9CI)

MF C24 H30 O12

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-4-carboxylic acid, 1,4a,5,6,7,7a-hexahydro-1,4a,5,6,7-pentahydroxy-7-methyl-, methyl ester, [1S-(1.alpha.,4a.alpha.,5.alpha.,6.alpha.,7.alpha.,7a.alpha.)]- (9CI)

MF C11 H16 O8

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

REGISTRY COPYRIGHT 2003 ACS on STN L3 33 ANSWERS

.beta.-D-Glucopyranoside, (1S,4aS,5S,6R,7R,7aS)-6-bromo-1,4a,5,6,7,7a-IN

hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

C15 H23 Br O10 MF

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):12

REGISTRY COPYRIGHT 2003 ACS on STN L3 33 ANSWERS

.beta.-D-Glucopyranoside, (1S,4aS,5R,6R,7R,7aR)-1,4a,5,6,7,7a-hexahydro-IN 4a,5,6,7-tetrahydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

C15 H24 O11 MF

- L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN
- IN .beta.-D-Glucopyranoside, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)
- MF C15 H24 O10
- CI COM

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN
- MF C9 H14 O5

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN Cyclopenta[c]pyran-4-carboxylic acid, 1-(.beta.-D-glucopyranosyloxy) 1,4a,5,6,7,7a-hexahydro-4a,5,6,7-tetrahydroxy-7-methyl-, methyl ester,
 (1S,4aR,5R,6S,7R,7aR)- (9CI)

MF C17 H26 O13

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-ribo-Hexopyranosid-3-ulose, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7ahexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl, 3-hydrate (9CI)

MF C15 H24 O11

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-Glucopyranoside, (1S,4aS,5S,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

MF C15 H24 O10

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

MF C27 H36 O17

CI IDS

CM 1

CM 2

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN
IN Cyclopenta[c]pyran-4-carboxylic acid, 1-(.beta.-D-glucopyranosyloxy)1,4a,5,6,7,7a-hexahydro-4a,5,6,7-tetrahydroxy-7-methyl-, methyl ester,
(1S,4aR,5R,6R,7R,7aR)- (9CI)
MF C17 H26 O13

$$HO-CH_2$$
 OH OH OH OH OH OH OH

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

.beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7methylcyclopenta[c]pyran-1-yl, 2,3,4,6-tetraacetate, [1S(1.alpha.,4a.alpha.,5.alpha.,7.alpha.,7a.alpha.)]- (9CI)

MF C23 H32 O14 CI COM

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

IN .beta.-D-ribo-Hexopyranosid-3-ulose, (1S,4aS,5R,7S,7aR)-1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-7-methylcyclopenta[c]pyran-1-yl (9CI)

MF C15 H22 O10

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN
IN .beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,7-trihydroxy-4-

(hydroxymethyl)-7-methylcyclopenta[c]pyran-1-yl (9CI)

MF C16 H26 O11

L3 33 ANSWERS REGISTRY COPYRIGHT 2003 ACS on STN

.beta.-D-Glucopyranoside, 1,4a,5,6,7,7a-hexahydro-4a,5,6,7-tetrahydroxy-7-methylcyclopenta[c]pyran-1-yl, heptaacetate, [1S-(1.alpha.,4a.alpha.,5.beta.,6.beta.,7.alpha.,7a.alpha.)]- (9CI) IN

MF C29 H38 O18

 \mathtt{CI} IDS

> CM1

Absolute stereochemistry.

CM 2

ALL ANSWERS HAVE BEEN SCANNED

=> s 13 and composition

7 COMPOSITION

2 COMPOSITIONS

9 COMPOSITION

(COMPOSITION OR COMPOSITIONS)

L7 0 L3 AND COMPOSITION

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(FILE 'HOME' ENTERED AT 10:53:49 ON 22 DEC 2003)

FILE 'REGISTRY' ENTERED AT 10:53:59 ON 22 DEC 2003

STRUCTURE UPLOADED

L2 2 S L1 SSS SAM

L3 33 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 10:55:20 ON 22 DEC 2003

4 S L3 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)

FILE 'REGISTRY' ENTERED AT 10:56:54 ON 22 DEC 2003

L5 1 S 6926-08-5

L6 0 S L5 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)

L7 0 S L3 AND COMPOSITION

=> file polymers

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plantlets from nodal cuttings and their acclimatization to ex vitro conditions. Single node cuttings were submitted to a root induction

treatment with .beta.-indoleacetic acid (5 days at 2 mg 1-1) followed by a transfer to a phytohormone-free medium to promote root elongation and support plantlet development. The new plantlets were weaned under autotrophic conditions and subsequently acclimatized in a glasshouse where they grew into fertile flowering plants that produced the characteristic Devil's Claw fruits as well as tuberised roots. Anal. of the tuber tissue of the micropropagated plants showed the presence of the iridoids harpagoside and harpagide at concns. comparable with those found in the wild plant material (1% dry wt.). The leaves were also found to contain these iridoids, and therefore could potentially provide an alternative and more sustainable source of therapeutically active compds. The application of in vitro methods for the propagation of Devil's Claw would contribute to the conservation of this species.

THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 15 ALL CITATIONS AVAILABLE IN THE RE FORMAT

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ANSWER 2 OF 11 CAPLUS COPYRIGHT 2003 ACS on STN
L9
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2002:591953 CAPLUS AN

DN 137:159305

TI 2-0-(9Z,12Z-octadecadienoyl)-3-0-[.alpha.-D-galactopyranosyl-(1''-6')-0-.beta.-D-galactopyranosyl]glycerol and pharmaceutical formulations containing it

IN Shin, Jun Shik; Kim, Sang Tae; Hahn, Yong Nam

PA Japan

SO Jpn. Kokai Tokkyo Koho, 30 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

| | PATENT NO. | | DATE | APPLICATION NO. | DATE | | |
|------|---------------|----|----------|-----------------|----------|--|--|
| | | | | | | | |
| PΙ | JP 2002220400 | A2 | 20020809 | JP 2001-365399 | 20011129 | | |
| | US 6531582 | B1 | 20030311 | US 2001-995617 | 20011129 | | |
| PRAI | KR 2000-71438 | A | 20001129 | | | | |

AB Pharmaceutical formulations for treatment of osteoporosis, arthritis, or intervertebral disk hernia, contain 2-0-(9Z,12Z-octadecadienoy1)-3-0-[.alpha.-D-galactopyranosy1-(1''-6')-0-.beta.-D-galactopyranosyl]glycerol (I) or its esters as active ingredients. I (420 mg) was purified from an EtOH ext. of 1848 g Cibotium barometz root powder. Administration of I at 75 .mu.g/mL p.o. for 2 wk prevented mouse paw edema induced by Zymosan A and Freund's adjuvant. Formulation examples of injections, tablets, capsules, and liqs. contq. I or I acetate are given.

- L9 ANSWER 3 OF 11 CAPLUS COPYRIGHT 2003 ACS on STN
- AN2002:533182 CAPLUS

DN 137:88448

ΤI Use of harpagide-related compounds for prevention and treatment of osteoporosis, arthritis, and intervertebral disk hernia, pharmaceutical compositions, and preparation of the compounds IN

Shin, Jun Sik; Kim, Sang-Tae; Hahn, Yong-Nam

PA S. Korea

SO Jpn. Kokai Tokkyo Koho, 29 pp.

CODEN: JKXXAF

- DTPatent
- LA Japanese

| FAN. | CNT 1 | | | | | | |
|------|------------------|------|----------|-----------------|----------|--|--|
| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE. | | |
| | | | | | | | |
| ΡI | JP 2002201136 | A2 | 20020716 | JP 2001-365400 | 20011129 | | |
| | US 2002183264 | A1 | 20021205 | US 2001-995691 | 20011129 | | |
| PRAI | KR 2000-71497 | A | 20001129 | | | | |
| OS | MARPAT 137:88448 | | | | | | |
| | | | | | | | |

GI

AB Harpagide-related compds. I (R1 = H, lower alkyl; R2 = H, cinnamoyl) are used for treatment or prevention of osteoporosis, arthritis, and/or intervertebral disk diseases. I (R1 = H, lower alkyl; R2 = cinnamoyl) are hydrolyzed to give I (R1 = H, lower alkyl; R2 = H). Harpagide (purified from Harpagophytum procumbens root) (at 75 .mu.g/kg/day p.o.) inhibited zymosan A- and Freund's adjuvant-induced rat paw edema. Formulation examples of injections, tablets, capsules, and liqs. contg. harpagide or harpagoside are given.

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ANSWER 4 OF 11 CAPLUS COPYRIGHT 2003 ACS on STN
1.9
AN
     2001:31340 CAPLUS
DN
     134:95502
     Compositions and methods for treating or preventing osteoporosis
TI
IN
     Prince, Richard Lewis; Min, Xu
     University of Western Australia, Australia; Guangzhou University of
     Traditional Chinese Medicine
SO
     PCT Int. Appl., 93 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
FAN.CNT 1
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| | PATENT | NO. | | KI | ND | DATE | | | A. | PPLI | CATI | ON N | 0.] | DATE | | | |
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| | | | | | | | | | _ | | | - - | | - | | | |
| PΙ | WO 2001 | 00199 | 96 | A | 1 | 2001 | 0111 | | W | 20 | 0 0 - A | U737 | : | 2000 | 0629 | | |
| | WO 2001 | 00199 | 96 | C: | 2 | 2002 | 0912 | | | | | | | | | | |
| | W: | ΑE, | AG, | AL, | AM, | AT, | AU, | ΑZ, | BA, | BB, | BG, | BR, | BY, | ΒZ, | CA, | CH, | CN, |
| | | CR, | CU, | CZ, | DE, | DK, | DM, | DΖ, | EE, | ES, | FI, | GB, | GD, | GE, | GH, | GM, | HR, |
| | | HŰ, | ID, | ΙL, | IN, | IS, | JP, | KE, | KG, | ΚP, | KR, | ΚZ, | LC, | LK, | LR, | LS, | LT, |
| | | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NO, | NZ, | PL, | PT, | RO, | RU, |
| | | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, | ΤZ, | UA, | UG, | US, | UΖ, | VN, |
| | | YU, | ZA, | ZW, | AM, | ΑZ, | BY, | KG, | KZ, | MD, | RU, | TJ, | TM | | | | |
| | RW: | GH, | GM, | ΚE, | LS, | MW, | MZ, | SD, | SL, | ŞΖ, | TZ, | UG, | ZW, | ΑT, | BE, | CH, | CY, |
| | | DE, | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | ΙT, | LU, | MC, | NL, | PT, | SE, | BF, | ВJ, |
| | | CF, | CG, | CI, | CM, | GA, | GN, | GW, | ML, | MR, | NE, | SN, | TD, | TG | | | |

PRAI AU 1999-1273 A 19990629

AB The invention relates to a therapeutic compn. and method for treating osteoporosis and other calcium, and/or estrogen related disorders. Examples are given for treating osteoporosis with exts. of plants such as Epimedium koreanum, Slavia miltiorrhiza, Asragalus membranaceus, Pueraria thomsonii, and Psoralea coryliofolia.

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L9 ANSWER 5 OF 11 IFIPAT COPYRIGHT 2003 IFI on STN AN 03842228 IFIPAT; IFIUDB; IFICDB
- TI 2-O-(9Z,12Z-OCTADECADIENOYL)-3-O-(ALPHA-D-GALACTOPYRANOSYL-(1''-6')-O-ALPHA -D-GALACTOPYRANOSYL)GLYCEROL AND PHARMACEUTICAL COMPOSITION CONTAINING THE SAME
- INF Han; Yong Nam, Seoul, KR
 Kim; Sang Tae, Seoul, KR
 Shin; Joon Shik, Hospital of Jaseng Oriental Medicine, 635, Shinsa-dong,
 Kangnam-ku, Seoul, KR
- IN Han Yong Nam (KR); Kim Sang Tae (KR); Shin Joon Shik (KR)
- PAF Shin; Joon Shik, Seoul, KR

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Unassigned Or Assigned To Individual (68000)
PA
EXNAM Henley, III, Raymond
     Birch Stewart Kolasch & Birch LLP
AG
                     B1 20030311
     US 6531582
PΙ
                          20011129
     US 2001-995617
ΑI
     29 Nov 2021
XPD
                          20001129
     KR 2000-71438
PRAI
                          20030311
     US 6531582
FI
     Utility
DT
      CHEMICAL
FS
      GRANTED
      012334
               MFN: 0870
MRN
CLMN
      17 Drawing Sheet(s), 27 Figure(s).
GT
     FIG. 1 shows (a) the flowchart of a process for recovering the dry weight
     of the pharmaceutical composition by extraction of the constituent drug
      with distilled water and an organic solvent, concentration under reduced
     pressure and lyophilization, and (b) the flowchart of the procedures for
      extracting the organic fraction according to the process for extracting
      the effective component.
     FIG. 2 shows (a) the causal mechanism of invasion, (b) the invaded
      portions, and (c) the route for transmitting molecular biological signals
      of invasive process in joint at which arthritis as the typical
      one of bone diseases is invaded.
     FIG. 3 shows the survival and cytomorphological appearance of synovial
      cells in joint portion at which arthritis as the typical
      chronic and degenerative bone diseases is invaded.
     FIG. 4 shows the 70-days inhibitory effect on edema as one of chronic and
      degenerative osteopathological symptoms of arthritis by
      treating the induced edema with respective fractions obtained as the
      organic solvent fractions of the pharmaceutical composition in an amount
      of 75 mu g/ml for 2 weeks via oral route.
     FIG. 5 shows the number and distribution pattern of CAM observed for 3
      days by treating the fertilized egg with the effective component as the
      organic solvent extract in the concentration of 10 mu g/ml, incubating
      the egg in an incubator at 37 degrees C. for 2 days and then carefully
      injecting 5 x 103 synovial cells on CAM via syringe.
     FIG. 6 is H/E tissue staining which shows the rupture extent of
      cartilaginous tissue after 70 days from the treatment of
      arthritis-induced animal with the water extract of the
      pharmaceutical composition at the concentration given in FIG. 9.
     FIG. 7 is (a) a bar graph showing the inhibition of NO formation when Raw
      264.7 cell lines are treated with the organic solvent fraction of a
      single drug of the pharmaceutical composition at the concentration of 10
      mu g/ml for 24 hours an then stimulated with LPS, and (b) a bar graph
      showing the inhibition of NO formation when the cell lines treated with
      CBB fraction and LNE fraction as the organic solvent fractions are
      stimulated with LPS according to the same manner as above (a).
     FIG. 8 shows (a) the induction pattern of apoptosis when Raw 264. 7 cell
      lines are treated with the organic fraction of the effective component at
      the concentration of about 20 mu g/ml and then stimulated with LPS, and
      (b) the result of observing whether the synovial cells show the same
      pattern according to the same method as above (a).
     FIG. 9 shows (a) the result of flow cytometry to determine the effect on
      cell cycle by treating Raw 264.7 cell lines with the organic fraction of
      the effective component at the concentration of about 20 mu g/ml and then
      stimulating with LPS and (b) the result of observing whether the synovial
      cells show the same appearance according to the same method as above (a).
     FIG. 10 shows the inhibitory effect on the expression of COX-II enzyme
      protein as measured by SDS-PAGE electrophoresis when synovial cell lines
      are treated with the organic fraction of the effective component and then
      stimulated with LPS.
     FIG. 11 shows the inhibitory effect on the synthesis of iNOS and COX-II
      enzymes as measured by RT-PCR when synovial cell lines are treated with
```

the organic fraction of the effective component and then stimulated with

LPS, and the inhibitory effect of respective fractions on the synthesis of iNOS (b) and COX-II (c) enzymes according to the same method as above (a).

FIG. 12 shows the result obtained by labeling synovial cell lines with a secondary antibody FITC, allowing to stand the cells for about one hour while shading the light with a foil and then observing the cells under a fluorescence microscope, in order to examine whether the compound identified as the effective component in the pharmaceutical composition of the present invention can induce the inhibitory effect on COX-II expression in synovial cells in joint portion.

FIG. 13 is X-ray photograph to show the 70-days inhibitory effect on edema as one of chronic and degenerative osteopathological symptoms of arthritis by treating the induced edema with respective fractions obtained as the organic solvent fractions of the pharmaceutical composition in an amount of 75 mu g/ml for 2 weeks via oral route.

FIG. 14 is the result of computerized tomography (CT) to show the clinical improvement in an outpatient suffering from ruptured **disc** with the pharmaceutical composition of the present invention.

FIG. 15 is the result of magnetic resonance imaging (MRI) to show the clinical improvement in an outpatient suffering from ruptured disc with the pharmaceutical composition of the present invention.

FIG. 16 shows the presence of nogo-A with respect to the mechanism to induce vertebral neuroparalysis in an outpatient suffering from ruptured disc.

FIG. 17 shows a channel for blocking neurotransmission by raising the injury in oligodendrocyte present around the axon as the nervous portion concerned with a paralysis of neurotransmission.

FIG. 18 shows (a) a channel for blocking neurotransmission to brain cells as in case that the injury is raised in oligodendrocyte present around the axon as the nervous portion concerned with a paralysis of neurotransmission, (b) the recovery of neurotransmission by treating cells with NGF or CBB13/LNE-2 to regenerate neutrite, which recovers the neurotransmission.

The present invention relates to the novel compound 2-0-(9z,12zoctadecadienoyl)-3-0-(alpha -D-galactopyranosyl-(1''-6')-Oalpha -D-galactopyranosyl)glycerol (Generic name: shinbarometin) having the chemical structure represented by the following formula:

DRAWING

or its acetate having an excellent effect on arthritis, osteoporosis and ruptured disc, and to a pharmaceutical composition containing said compound as an effective component, in combination with a pharmaceutically acceptable auxiliary, diluent, isotonic agent, preservative, lubricant and solubilizing aid, which is formulated in the form of a pharmaceutically acceptable preparation and has a potent effect for osteoporosis, arthritis and ruptured disc.

CLMN 3

AΒ

- GI 17 Drawing Sheet(s), 27 Figure(s).
 - FIG. 1 shows (a) the flowchart of a process for recovering the dry weight of the pharmaceutical composition by extraction of the constituent drug with distilled water and an organic solvent, concentration under reduced pressure and lyophilization, and (b) the flowchart of the procedures for extracting the organic fraction according to the process for extracting the effective component.
 - FIG. 2 shows (a) the causal mechanism of invasion, (b) the invaded portions, and (c) the route for transmitting molecular biological signals of invasive process in joint at which **arthritis** as the typical one of bone diseases is invaded.
 - FIG. 3 shows the survival and cytomorphological appearance of synovial cells in joint portion at which **arthritis** as the typical chronic and degenerative bone diseases is invaded.
 - FIG. 4 shows the 70-days inhibitory effect on edema as one of chronic and

degenerative osteopathological symptoms of **arthritis** by treating the induced edema with respective fractions obtained as the organic solvent fractions of the pharmaceutical composition in an amount of 75 mu g/ml for 2 weeks via oral route.

- FIG. 5 shows the number and distribution pattern of CAM observed for 3 days by treating the fertilized egg with the effective component as the organic solvent extract in the concentration of 10 mu g/ml, incubating the egg in an incubator at 37 degrees C. for 2 days and then carefully injecting 5 x 103 synovial cells on CAM via syringe.
- FIG. 6 is H/E tissue staining which shows the rupture extent of cartilaginous tissue after 70 days from the treatment of arthritis-induced animal with the water extract of the pharmaceutical composition at the concentration given in FIG. 9.
- FIG. 7 is (a) a bar graph showing the inhibition of NO formation when Raw 264.7 cell lines are treated with the organic solvent fraction of a single drug of the pharmaceutical composition at the concentration of 10 mu g/ml for 24 hours an then stimulated with LPS, and (b) a bar graph showing the inhibition of NO formation when the cell lines treated with CBB fraction and LNE fraction as the organic solvent fractions are stimulated with LPS according to the same manner as above (a).
- FIG. 8 shows (a) the induction pattern of apoptosis when Raw 264. 7 cell lines are treated with the organic fraction of the effective component at the concentration of about 20 mu g/ml and then stimulated with LPS, and (b) the result of observing whether the synovial cells show the same pattern according to the same method as above (a).
- FIG. 9 shows (a) the result of flow cytometry to determine the effect on cell cycle by treating Raw 264.7 cell lines with the organic fraction of the effective component at the concentration of about 20 mu g/ml and then stimulating with LPS and (b) the result of observing whether the synovial cells show the same appearance according to the same method as above (a).
- FIG. 10 shows the inhibitory effect on the expression of COX-II enzyme protein as measured by SDS-PAGE electrophoresis when synovial cell lines are treated with the organic fraction of the effective component and then stimulated with LPS.
- FIG. 11 shows the inhibitory effect on the synthesis of iNOS and COX-II enzymes as measured by RT-PCR when synovial cell lines are treated with the organic fraction of the effective component and then stimulated with LPS, and the inhibitory effect of respective fractions on the synthesis of iNOS (b) and COX-II (c) enzymes according to the same method as above (a).
- FIG. 12 shows the result obtained by labeling synovial cell lines with a secondary antibody FITC, allowing to stand the cells for about one hour while shading the light with a foil and then observing the cells under a fluorescence microscope, in order to examine whether the compound identified as the effective component in the pharmaceutical composition of the present invention can induce the inhibitory effect on COX-II expression in synovial cells in joint portion.
- FIG. 13 is X-ray photograph to show the 70-days inhibitory effect on edema as one of chronic and degenerative osteopathological symptoms of arthritis by treating the induced edema with respective fractions obtained as the organic solvent fractions of the pharmaceutical composition in an amount of 75 mu g/ml for 2 weeks via oral route.
- FIG. 14 is the result of computerized tomography (CT) to show the clinical improvement in an outpatient suffering from ruptured **disc** with the pharmaceutical composition of the present invention.
- FIG. 15 is the result of magnetic resonance imaging (MRI) to show the clinical improvement in an outpatient suffering from ruptured disc with the pharmaceutical composition of the present invention.
- FIG. 16 shows the presence of nogo-A with respect to the mechanism to induce vertebral neuroparalysis in an outpatient suffering from ruptured disc.
- FIG. 17 shows a channel for blocking neurotransmission by raising the injury in oligodendrocyte present around the axon as the nervous portion concerned with a paralysis of neurotransmission.

FIG. 18 shows (a) a channel for blocking neurotransmission to brain cells as in case that the injury is raised in oligodendrocyte present around the axon as the nervous portion concerned with a paralysis of neurotransmission, (b) the recovery of neurotransmission by treating cells with NGF or CBB13/LNE-2 to regenerate neutrite, which recovers the neurotransmission.

- L9 ANSWER 6 OF 11 PASCAL COPYRIGHT 2003 INIST-CNRS. ALL RIGHTS RESERVED. on STN
- AN 2002-0594962 PASCAL
- CP Copyright .COPYRGT. 2002 INIST-CNRS. All rights reserved.
- TIEN In vitro propagation and iridoid analysis of the medicinal species Harpagophytum procumbens and H. zeyheri
- AU LEVIEILLE G.; WILSON G.
- CS Department of Botany, University College Dublin, Belfield, Dublin, Ireland
- SO Plant cell reports : (Print), (2002), 21(3), 220-225, 15 refs. ISSN: 0721-7714 CODEN: PCRPD8
- DT Journal
- BL Analytic
- CY Germany, Federal Republic of
- LA English
- AV INIST-18737, 354000106592000060
- CP Copyright .COPYRGT. 2002 INIST-CNRS. All rights reserved.
- Extracts of the tubers of Harpagophytum proc-umbens DC (Devil's Claw) are AB used widely for the relief of arthritis, lumbago and muscular pain. The anti-inflammatory activity has been attributed to their iridoid components. A two-step protocol was established for the in vitro propagation of plants of Harpagophytum sp. by the regeneration of new plantlets from nodal cuttings and their acclimatisation to ex vitro conditions. Single node cuttings were submitted to a root induction treatment with .beta.-indoleacetic acid (5 days at 2 mg l.sup.-.sup.1) followed by a transfer to a phytohormone-free medium to promote root elongation and support plantlet development. The new plantlets were weaned under autotrophic conditions and subsequently acclimatised in a glasshouse where they grew into fertile flowering plants that produced the characteristic Devil's Claw fruits as well as tuberised roots. Analysis of the tuber tissue of the micropropagated plants showed the presence of the iridoids harpagoside and harpagide at concentrations comparable with those found in the wild plant material (1% dry weight). The leaves were also found to contain these iridoids, and therefore could potentially provide an alternative and more sustainable source of therapeutically active compounds. The application of in vitro methods for the propagation of Devil's Claw would contribute to the conservation of this species.
- L9 ANSWER 7 OF 11 SCISEARCH COPYRIGHT 2003 THOMSON ISI on STN
- AN 2002:937703 SCISEARCH
- GA The Genuine Article (R) Number: 614RL
- TI In vitro propagation and iridoid analysis of the medicinal species Harpagophytum procumbens and H-zeyheri
- AU Levieille G (Reprint); Wilson G
- CS Univ Coll Dublin, Dept Bot, Dublin 4, Ireland (Reprint)
- CYA Ireland
- PLANT CELL REPORTS, (OCT 2002) Vol. 21, No. 3, pp. 220-225.
 Publisher: SPRINGER-VERLAG, 175 FIFTH AVE, NEW YORK, NY 10010 USA.
 ISSN: 0721-7714.
- DT Article; Journal
- LA English
- REC Reference Count: 15
 - *ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS*
- AB Extracts of the tubers of Harpagophytum procumberzs DC (Devil's Claw) are used widely for the relief of **arthritis**, lumbago and muscular pain. The anti-inflammatory activity has been attributed to their iridoid components. A two-step protocol was established for the in vitro

propagation of plants of Harpagophytum sp. by the regeneration of new plantlets from nodal cuttings and their acclimatisation to ex vitro conditions. Single node cuttings were submitted to a root induction treatment with beta-indoleacetic acid (5 days at 2 mg l(-1)) followed by a transfer to a phytohormone-free medium to promote root elongation and support plantlet development. The new plantlets were weaned under autotrophic conditions and subsequently acclimatised in a glasshouse where they grew into fertile flowering plants that produced the characteristic Devil's Claw fruits as well as tuberised roots. Analysis of the tuber tissue of the micropropagated plants showed the presence of the iridoids harpagoside and harpagide at concentrations comparable with those found in the wild plant material (1% dry weight). The leaves were also found to contain these iridoids, and therefore could potentially provide an alternative and more sustainable source of therapeutically active compounds. The application of in vitro methods for the propagation of Devil's Claw would contribute to the conservation of this species.

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ANSWER 8 OF 11 USPATFULL on STN
L9
       2003:67836 USPATFULL
AN
       2-0-(9z,12z-octadecadienoyl)-3-0-[.alpha.-D-galactopyranosyl-(1"-6')-0-
ΤI
       .alpha.-D-galactopyranosyl]glycerol and pharmaceutical composition
       containing the same
       Shin, Joon Shik, Hospital of Jaseng Oriental Medicine, 635, Shinsa-dong,
TN
       Kangnam-ku, Seoul, KOREA, REPUBLIC OF
       Kim, Sang Tae, Seoul, KOREA, REPUBLIC OF Han, Yong Nam, Seoul, KOREA, REPUBLIC OF
       Shin, Joon Shik, Seoul, KOREA, REPUBLIC OF (non-U.S. individual)
PΑ
                                20030311
PΙ
       US 6531582
       US 2001-995617
                                20011129 (9)
ΑI
PRAI
       KR 2000-71438
                           20001129
DT
       Utility
FS
       GRANTED
       Primary Examiner: Henley, III, Raymond
EXNAM
       Birch Stewart Kolasch & Birch LLP
LREP
CLMN
       Number of Claims: 3
ECL
       Exemplary Claim: 1
       27 Drawing Figure(s); 17 Drawing Page(s)
DRWN
LN.CNT 971
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to the novel compound 2-0-(9z,12z-
       octadecadienoyl)-3-0-[.alpha.-D-galactopyranosyl-(1"-6')-0-.alpha.-D-
       galactopyranosyl]glycerol (Generic name: shinbarometin) having the
       chemical structure represented by the following formula: ##STR1##
       or its acetate having an excellent effect on arthritis,
       osteoporosis and ruptured disc, and to a
       pharmaceutical composition containing said compound as an effective
       component, in combination with a pharmaceutically acceptable auxiliary,
       diluent, isotonic agent, preservative, lubricant and solubilizing aid,
       which is formulated in the form of a pharmaceutically acceptable
       preparation and has a potent effect for osteoporosis,
       arthritis and ruptured disc.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     ANSWER 9 OF 11 USPATFULL on STN
L9
       2002:323098 USPATFULL
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Use of harpagid-related compounds for prevention and treatment of

osteoporosis, arthritis and ruptured disc

Shin, Joon Shik, Seoul, KOREA, REPUBLIC OF

Kim, Sang Tae, Seoul, KOREA, REPUBLIC OF Han, Yong Nam, Seoul, KOREA, REPUBLIC OF

A1

A1

and pharmaceutical composition containing the same

20021205

20011129 (9)

AN

TТ

IN

PΙ

ΑI

US 2002183264

US 2001-995691

KR 2000-71497 20001129 PRAI

DТ Utility

FS APPLICATION

BIRCH STEWART KOLASCH & BIRCH, PO BOX 747, FALLS CHURCH, VA, 22040-0747 LREP

CLMN Number of Claims: 4 ECL Exemplary Claim: 1 DRWN 17 Drawing Page(s)

LN.CNT 1078

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

In the present invention, it is discovered that a compound of formula (I) has a potent effect for treatment of osteoporosis, arthritis and ruptured disc: ##STR1##

in which R.sub.1 represents hydrogen atom or alkyl group and R.sub.2 represents hydrogen atom r cinnamoyl group. Therefore, the compound of formula (I) can be used for prevention and treatment of osteoporosis, arthritis and ruptured disc. Thus, the present invention provides a pharmaceutical preparation containing as an effective component a compound of formula (I) in combination with a pharmaceutically acceptable auxiliary, diluent, isotonic agent, preservative, lubricant and solubilizing aid, which is formulated in the form of a pharmaceutically acceptable preparation and has a potent effect for osteoporosis, arthritis and

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Ь9 ANSWER 10 OF 11 USPATFULL on STN

AN 2001:141886 USPATFULL

ruptured disc.

TIHarpagoside-enriched extract from harpagophytum procumbens and processes for producing same

Stumpf, Karl-Heinz, Karlsruhe, Germany, Federal Republic of Jaggy, Hermann, Karlsruhe, Germany, Federal Republic of Oschmann, Rainer, Landau, Germany, Federal Republic of Koch, Egon, Karlsruhe, Germany, Federal Republic of Simmet, Thomas, Bochum, Germany, Federal Republic of

Dr. Willmar Schwabe GmbH & Co., Karlsruhe, Germany, Federal Republic of PA

(non-U.S. corporation)

PΙ US 6280737 20010828

WO 9734565 19970925

US 1999-155043 19990222 (9) AΤ

WO 1997~DE591 19970321

> 19990222 PCT 371 date 19990222 PCT 102(e) date

PRAI DE 1996-19611221 19960321

DE 1996-19651290 19961210

DΤ Utility

IN

FS GRANTED

EXNAM Primary Examiner: Lilling, Herbert J. LREP McDonnell Boehnen Hulbert & Berghoff

CLMN Number of Claims: 9

ECL Exemplary Claim: 1

DRWN No Drawings LN.CNT 313

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The disclosure relates to extracts from Harpagophytum procumbens with a high harpagoside content, to processes for producing them, such extracts containing no components capable of stimulating the synthesis of thromboxane B.sub.2 and cysteinylleucotrienes, and to pharmaceutical compositions containing such extracts.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 11 OF 11 WPINDEX COPYRIGHT 2003 THOMSON DERWENT on STN

AN 2003-010740 [01] WPINDEX

```
DNC C2003-002594
     A new compound, 2-0-(9z,12z-octadecadienyl)-3-0-(alpha-galactopyranosyl-
TΙ
     (1'-6')-O-beta-D-galactopyranosyl)glycerol, useful for the treatment of
     osteoporosis, arthritis, and extruded discs.
DC
     HAN, Y N; KIM, S T; SHIN, J S
IN
     (SHIN-I) SHIN S; (SHIN-I) SHIN J S
PA
CYC
    3
     KR 2002041837 A 20020605 (200301)*
JP 2002220400 A 20020809 (200305)
                                                1p
ΡI
                                               g08
     US 6531582 B1 20030311 (200321)
   KR 2002041837 A KR 2000-71438 20001129; JP 2002220400 A JP 2001-365399
     20011129; US 6531582 B1 US 2001-995617 20011129
                      20001129
PRAI KR 2000-71438
     2003-010740 [01]
                        WPINDEX
     KR2002041837 A UPAB: 20030101
AB
     NOVELTY - Provided are novel 2-0-(9z,12z-octadecadienyl)-3-0-( alpha
     -galactopyranosyl-(1'-6')-O- beta -D-galactopyranosyl)glycerol (I) having
     excellent effects on osteoporosis, arthritis, and
     extruded disc.
          DETAILED DESCRIPTION - 2-0-(9z,12z-octadecadienyl)-3-0-( alpha
     -qalactopyranosyl-(1'-6')-0- beta -D-galactopyranosyl)glycerol is
     represented by the following structural formula.
          An INDEPENDENT CLAIM is also included for a pharmaceutical
     preparation containing (I) it or its acetate and one of allendrate,
     tamoxifen, vitamin B3, parathyroid hormone, sulfasalazine, thioredoxin
     reductase, alendronate, raloxifene, calcitonin, estradiol, genistein,
     1,25-dihydroxyvitamin D3, alendronate, estrogen receptor modulator,
     biphosphonates and harpagide, as active ingredients, and
     pharmaceutically acceptable excipients, diluents, adjuvants,
     preservatives, dissolution adjuvants.
     Dwg.1/10
=> dis hist
     (FILE 'HOME' ENTERED AT 10:53:49 ON 22 DEC 2003)
     FILE 'REGISTRY' ENTERED AT 10:53:59 ON 22 DEC 2003
L1
                STRUCTURE UPLOADED
              2 S L1 SSS SAM
L2
             33 S L1 SSS FULL
L3
     FILE 'CAPLUS' ENTERED AT 10:55:20 ON 22 DEC 2003
              4 S L3 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)
L4
     FILE 'REGISTRY' ENTERED AT 10:56:54 ON 22 DEC 2003
              1 S 6926-08-5
L5
              0 S L5 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)
L6
              0 S L3 AND COMPOSITION
L7
     FILE 'APOLLIT, BABS, CAPLUS, CBNB, CEN, CIN, DISSABS, EMA, IFIPAT,
     JICST-EPLUS, PASCAL, PLASNEWS, PROMT, RAPRA, SCISEARCH, TEXTILETECH,
     USPATFULL, USPAT2, WPINDEX, WTEXTILES' ENTERED AT 11:03:18 ON 22 DEC 2003
            249 S HARPAGIDE
L8
             11 S L8 AND (ARTHRITIS OR OSTEOPOROSIS OR DISC)
L9
=>
---Logging off of STN---
```

Executing the logoff script...

=> LOG Y

| COST IN U.S. DOLLARS | SINCE FILE
ENTRY | TOTAL
SESSION |
|--|---------------------|------------------|
| FULL ESTIMATED COST | 63.54 | 258.14 |
| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE
ENTRY | TOTAL
SESSION |
| CA SUBSCRIBER PRICE | -2.60 | -5.20 |

STN INTERNATIONAL LOGOFF AT 11:05:22 ON 22 DEC 2003